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Article

A series on intestinal strongyloidiasis in immunocompetent and immunocompromised hosts

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ABSTRACT

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Background: Strongyloidiasis, endemic in tropical areas, may be asymptomatic in immunocompetent subjects or may cause potentially fatal hyper-infection in immunocompromised patients.

Methods: Of the 13,885 patients referred to the parasitology laboratory at our tertiary care referral center for stool microscopy, 15 were diagnosed as strongyloidiasis over a 6 year period. We assessed these patients retrospectively.

Results: Most patients were young (median age 32 years, range 3-66) males (12, 80%). Seven patients (46.6%) were immunocompromised. All patients were symptomatic, and symptoms included chronic diarrhea (4, 26.7%), acute diarrhea (1, 6.7%), abdominal pain (6, 40%), weight loss (3, 20%), cough (2, 13.33%), vomiting (1, 6.7%), anemia (10, 66.7%) and eosinophilia (3, 20%). Thirteen patients (86.6%) were diagnosed on first stool microscopy. Duodenal biopsy showed normal histology in twelve (80%) and partial villous atrophy in one (6.7%) patient. Stool microscopy also revealed giardiasis and cryptosporidiosis in one patient each. Nine patients responded well to ivermectin and albendazole, one died and five were lost to follow-up.

Conclusions: In endemic areas, even immunocompetent subjects may suffer from symptomatic strongyloidiasis and associated eosinophilia is uncommon.

KEYWORDS: chronic diarrhea, villous atrophy, parasite, eosinophilia

Introduction

Strongyloides stercoralis is a nematode endemic in tropical and subtropical areas.¹ Infected individuals may remain asymptomatic, but fatal illness or disseminated strongyloidiasis may develop in immunocompromised hosts. Strongyloidiasis may present with indigestion, abdominal pain, vomiting, chronic diarrhea, protein losing enteropathy and weight loss. Since symptoms of strongyloidiasis are often non-specific, it is rarely suspected or diagnosed clinically. Strongyloidiasis may be suspected by the clinicians in presence of peripheral blood eosinophilia. However, eosinophilia may not occur in

hyper-infection, particularly in immunocompromised hosts. Immunocompromised states associated with strongyloidiasis include acquired immunodeficiency syndrome, chronic corticosteroid treatment, and organ transplantation such as kidney, liver, bone marrow transplantation, as these patients receive multiple immunosuppressive drugs. Hence, we performed a retrospective analysis on 15 consecutive patients with *Strongyloides stercoralis* infestation diagnosed by stool microscopy in the Department of Microbiology of our tertiary care referral center with the aim to evaluate: (i) frequency of

immunocompetent or immunocompromised states among patients with strongyloidiasis, and (b) peripheral blood eosinophilia among patients with strongyloidiasis with or without immunocompromised states.

Methods

Data collection

All patients diagnosed with strongyloidiasis on stool microscopy at our parasitology laboratory over a 6 year period from April 2004 to July 2010, were included in this study. The demographic, clinical and laboratory data of the patients were collected from the electronic hospital information system and/or case records.

Laboratory procedures

Each patient underwent stool microscopy using direct wet mount technique. An attempt was made to examine three consecutive stool specimens until the diagnosis of intestinal strongyloidiasis could be confirmed. Normal saline and iodine preparation were used for stool examination. Peripheral blood was examined for total and differential leukocyte count; an absolute eosinophil count greater than 400 cells/mm³ was considered as eosinophilia. Disseminated infection was diagnosed from case records if a patient had evidence of extra-intestinal involvement.

Treatment and follow-up

All strongyloidiasis patients were treated either with ivermectin (6 mg twice daily for 6 days in an immunocompromised host and for five days in an immunocompetent host) or albendazole (400 mg twice daily for 3 days). The follow-up data of patients were retrieved from hospital records. Stool microscopy was repeated on follow-up in patients consenting for it.

Data analysis

The data collected were analyzed using SPSS version 15.0.

Results

Demographic and clinical parameters

15/13,885 (0.1%) patients who underwent stool microscopy at our laboratory were found to have strongyloidiasis. Most

patients were young (median age 32 years, range 3 to 66) males (12/15, 80%).

Of fifteen patients, 7 (46.6%) were immunocompromised; 6 were on corticosteroids and one was infected with human immunodeficiency virus. All the patients were symptomatic and the symptoms included chronic diarrhea (>4 weeks duration) in four (26.7%), acute diarrhea in one (6.7%), abdominal pain in six (40%), weight loss in three (20%), cough in two (13.3%) and vomiting in one (6.7%). Ten patients (66.7%) had anemia and three (20%) had eosinophilia.

Laboratory parameters

Laboratory parameters are summarized in **Table 1** and **Table 2**. Strongyloidiasis was diagnosed on first stool microscopy in most of the patients (13, 86.6%). Stool microscopy revealed ova of *Strongyloides* in a patient who also had acute renal

Table 1: Laboratory parameters of the patients

Variables	Immunocompetent patients (n=8)	Immunocompromised patients (n=7)	p value	Normal
Clinical variables (Median and range)				
Age (years)	41.50 (14-52)	22 (3-66)	0.37	
Laboratory variable (Median and range)				
Hemoglobin (g/dL)	11.6 (7.2–13.8)	9 (7.2-15.7)	0.45	12-16
TLC (cells/ μ L)	8250 (6000–19000)	7800 (2200-17600)	0.91	4000-11000
Neutrophil (cells/ μ L)	5497 (1826-8550)	3520 (2002-12060)	0.73	2000-7500
Lymphocyte (cells/ μ L)	2483.5 (1080-6225)	1312 (176-13904)	0.47	1500-4000
Eosinophil (cells/mm ³)	254 (0-6650)	82 (0-1296)	0.06	0-400
Eosinophilia*	2	1	0.83	-
Monocyte (cells/ μ L)	145 (0-537)	156 (22-704)	0.9	200-800
ESR (mm/hr)	20 (5-72)	20 (2-66)	0.93	0-15

*Absolute eosinophil count > 400 cells/mm³

Table 2: Comparison of clinical and laboratory parameters among immunocompetent and compromised patients

S.No.	Age/Gender	Clinical Diagnosis	Hb	TLC(\times 1000 cells/ μ L)	Eosinophil (cells/mm ³)
1	24/Male	Acute severe ulcerative colitis	7.7	4.1	82
2	54/Male	Symptomatic cholelithiasis	13.4	8.2	328
3	3/Male	Sepsis, hepatosplenomegaly	8.6	17.6	176
4	21/Male	Nephrotic syndrome	7.2	13.4	0
5	42/Female	Cushing's disease, pituitary macroadenoma, DM	10.1	11.4	114
6	52/Male	Lumb adenoma	7.2	6.2	372
7	28/Male	Reactive arthritis	11.5	17.9	6265
8	48/Female	Malena	11.8	7.1	0
9	32/Male	Chronic diarrhoea/ Chronic HBV	9.5	6	180
10	41/Male	Strongyloidiasis	13.8	19	6650
11	66/Male	multiple colonic ulcer	9	7.2	1296
12	6/Female	Chronic diarrhea	11	7.8	78
13	49/Male	Sensorymotor quadriplegia	15.7	9.4	188
14	22/Male	HIV, recurrent diarrhoea	9.6	2.2	22
15	14/Male	Solitary rectal ulcer	11.7	8.3	83

failure (**Figure 1 and 2**). Ova were of large size and had fully developed larvae. We could not do the molecular characterization to confirm species identification of these ova. Duodenal biopsy showed normal histology in twelve (80%), and partial villous atrophy in one (6.7%) patient. Duodenal biopsy revealed larvae of *Strongyloides* in one patient. Enzyme linked immunosorbent assay (ELISA) for human immunodeficiency virus was positive in one patient. *Giardia* and *Cryptosporidium* were detected on stool microscopy in one patient each. Three patients had eosinophilia, of which, two were immunocompetent and one was immunocompromised. *Strongyloides* hyper-infection occurs in immunosuppressive states due to disease or therapy with corticosteroids and is characterized by invasion of extra-intestinal sites like, lungs, liver, spleen, pancreas, gall bladder, kidneys, thyroid, brain, skin, and skeletal muscle. None of the patients had *Strongyloides* hyper-infection.



Figure 1: Lugol's Iodine stain of stool sample of a patient showing larva of *Strongyloides stercoralis*



Figure 2: Lugol's Iodine stain of stool sample of a patient showing hatching stage of *Strongyloides*

Treatment and follow-up

Fourteen patients were treated with ivermectin (6 mg twice daily for 6 days in immunocompromised host and for 5 days in an immunocompetent host) and one with albendazole (400 mg twice daily for 3 days). Nine patients showed resolution of infection and disappearance of *Strongyloides* over a median follow-up period of 14 days (range 4-570), one died within five days after diagnosis and five were lost to follow-up.

Discussion

The present retrospective study showed that, (a) in endemic areas, even immunocompetent subjects may suffer from symptomatic strongyloidiasis, (b) eosinophilia is uncommon in strongyloidiasis, and (c) first stool examination detects the parasite in most patients with strongyloidiasis on properly conducted stool microscopy.

Strongyloidiasis, endemic in tropical and subtropical countries, is usually known to be asymptomatic. However, it can cause fatal hyper-infection in patients with immunocompromised status. In a study on 78 asymptomatic individuals from southern India, the prevalence rate of parasitic infections was reported to be 97.4%, with *Strongyloides* infection rate of 20%.² Similarly, in another community based study of 198 individuals from Assam, 17 (8.5%) individuals were infected with *Strongyloides*, of which 10 (58.8%) were reported to be asymptomatic.³ In our study, all patients were symptomatic and presented with chronic diarrhea, acute diarrhea, abdominal pain, weight loss, cough, vomiting and anemia. Similarly, in a study describing five patients with strongyloidiasis in an endemic area, all presented with gastrointestinal symptoms.⁴ In another study on 205 parasitic infected patients, 68 (33.2%) had strongyloidiasis, of which 72% reported gastrointestinal symptoms.⁵ In addition to the above documentations there are other reports of patients infected with strongyloidiasis presenting with gastrointestinal symptoms.⁶⁻⁸ This suggests that clinicians should consider investigating for *Strongyloides stercoralis* infection even in immunocompetent patients presenting with suggestive gastrointestinal symptoms.

Strongyloides infection is known to be associated with peripheral blood eosinophilia in the immunocompetent host. However, this is considered unreliable in patients receiving corticosteroid therapy or those infected with human

immunodeficiency virus (HIV). In an earlier report of 12% patients infected with strongyloidiasis, 66% presented with eosinophilia.⁹ There are other reports suggesting eosinophilia to be an important laboratory finding in patients infected with *Strongyloides*.^{8,10–11} In contrast, we found eosinophilia to be rather uncommon even in immunocompetent individuals in our cohort. Absence of eosinophilia in immunocompromised patients diagnosed with strongyloidiasis has been reported in other studies.^{12–14} Hence, eosinophilia cannot be considered a reliable screening parameter for *Strongyloides* infection even in immunocompetent patients.

A single stool microscopy has been reported to be 30% sensitive for the diagnosis of intestinal *Strongyloides* infection.¹⁵ However, in our study 86.6% of the patients were diagnosed with strongyloidiasis on their first stool microscopy itself. This is similar to the finding of Grove et al¹⁶ who reported successful diagnosis of 68% patients with strongyloidiasis at the first stool microscopic examination. Furthermore, single stool microscopic examination has also been reported to reveal strongyloidiasis in a patient infected with AIDS.¹⁷ Therefore, first stool microscopic examination can be considered as an efficient technique for diagnosing strongyloidiasis.

The present study showed that most patients with *Strongyloides* infection to be male. The association was similar to that noted in previous studies.^{18–21} This high infection rate in males could be due to their outdoor work and hence higher probability of exposure to *Strongyloides stercoralis* infective stages.¹⁹ Another finding of the study was a higher *Strongyloides* infection rate in older age groups which was in concordance with findings of previous studies.¹⁹ This could be explained by the higher re-infection rates in the elderly.

We can conclude that, even immunocompetent individuals presenting with gastrointestinal symptoms should be considered for diagnosing *Strongyloides* infection even in the absence of eosinophilia. Detection of *Strongyloides* larva by direct stool microscopy is an efficient tool for diagnosing strongyloidiasis.

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